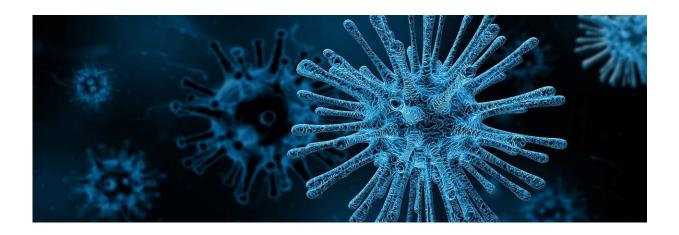


Another monkey virus could be poised for spillover to humans

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An obscure family of viruses, already endemic in wild African primates and known to cause fatal Ebola-like symptoms in some monkeys, is "poised for spillover" to humans, according to new University of Colorado Boulder research published online Sept. 30 in the journal *Cell*.

While such arteriviruses are already considered a critical threat to <u>macaque monkeys</u>, no <u>human infections</u> have been reported to date. And it is uncertain what impact the virus would have on people should it jump species.

But the authors, evoking parallels to HIV (the precursor of which



originated in African monkeys), are calling for vigilance nonetheless: By watching for arteriviruses now, in both animals and humans, the global health community could potentially avoid another pandemic, they said.

"This animal virus has figured out how to gain access to <u>human</u> cells, multiply itself, and escape some of the important immune mechanisms we would expect to protect us from an animal virus. That's pretty rare," said senior author Sara Sawyer, a professor of molecular, cellular and developmental biology at CU Boulder. "We should be paying attention to it."

There are thousands of unique viruses circulating among animals around the globe, most of them causing no symptoms. In recent decades, increasing numbers have jumped to humans, wreaking havoc on naïve immune systems with no experience fighting them off: That includes Middle Eastern Respiratory Syndrome (MERS) in 2012, Severe Acute Respiratory Syndrome coronavirus (SARS-CoV) in 2003, and SARS-CoV-2 (the virus that causes COVID-19) in 2020.

For 15 years, Sawyer's lab has used laboratory techniques and tissue samples from wildlife from around the globe to explore which animal viruses may be prone to jump to humans.

For the latest study, she and first author Cody Warren, then a postdoctoral fellow at the BioFrontiers Institute at CU, zeroed in on arteriviruses, which are common among pigs and horses but understudied among nonhuman primates. They looked specifically at simian hemorrhagic fever virus (SHFV), which causes a lethal disease similar to Ebola virus disease and has caused deadly outbreaks in captive macaque colonies dating back to the 1960s.

The study demonstrates that a molecule, or receptor, called CD163, plays a key role in the biology of simian arteriviruses, enabling the virus



to invade and cause infection of target cells. Through a series of laboratory experiments, the researchers discovered, to their surprise, that the virus was also remarkably adept at latching on to the <u>human</u> version of CD163, getting inside <u>human cells</u> and swiftly making copies of itself.

Like <u>human immunodeficiency virus</u> (HIV) and its precursor simian immunodeficiency virus (SIV), simian arteriviruses also appear to attack immune cells, disabling key defense mechanisms and taking hold in the body long-term.

"The similarities are profound between this <u>virus</u> and the simian viruses that gave rise to the HIV pandemic," said Warren, now an assistant professor in the College of Veterinary Medicine at The Ohio State University.

The authors stress that another pandemic is not imminent, and the public need not be alarmed.

But they do suggest that the global health community prioritize further study of simian arteriviruses, develop blood antibody tests for them, and consider surveillance of human populations with close contact to animal carriers.

A broad range of African monkeys already carries high viral loads of diverse arteriviruses, often without symptoms, and some species interact frequently with humans and are known to bite and scratch people.

"Just because we haven't diagnosed a human arterivirus infection yet doesn't mean that no human has been exposed. We haven't been looking," said Warren.

Warren and Sawyer note that in the 1970s, no one had heard of HIV



either.

Researchers now know that HIV likely originated from SIVs infecting nonhuman primates in Africa, likely jumping to humans sometime in the early 1900s.

When it began killing young men in the 1980s in the United States, no serology test existed, and no treatments were in the works.

Sawyer said there is no guarantee that these simian arteriviruses will jump to humans. But one thing is for sure: More viruses will jump to humans, and they will cause disease.

"COVID is just the latest in a long string of spillover events from animals to humans, some of which have erupted into global catastrophes," Sawyer said. "Our hope is that by raising awareness of the viruses that we should be looking out for, we can get ahead of this so that if human infections begin to occur, we're on it quickly."

More information: Sara L. Sawyer, Primate hemorrhagic fevercausing arteriviruses are poised for spillover to humans, *Cell* (2022). DOI: 10.1016/j.cell.2022.09.022. www.cell.com/cell/fulltext/S0092-8674(22)01194-1

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